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Please find below and/or attached an Office communication concerning this application or proceeding.

Office Action Summary

Application No. 09/922,426 Applicant(s)

Examiner

Maurie G. Baker, Ph.D.

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Trulson et al



-- The MAILING DATE of this communication appears on the cover sheet with the correspondence address --Period for Reply A SHORTENED STATUTORY PERIOD FOR REPLY IS SET TO EXPIRE THREE MONTH(S) FROM THE MAILING DATE OF THIS COMMUNICATION. - Extensions of time may be available under the provisions of 37 CFR 1.136 (a). In no event, however, may a reply be timely filed after SIX (6) MONTHS from the mailing date of this communication. - If the period for reply specified above is less than thirty (30) days, a reply within the statutory minimum of thirty (30) days will be considered timely. - If NO period for reply is specified above, the maximum statutory period will apply and will expire SIX (6) MONTHS from the mailing date of this communication. - Failure to reply within the set or extended period for reply will, by statute, cause the application to become ABANDONED (35 U.S.C. § 133). - Any reply received by the Office later than three months after the mailing date of this communication, even if timely filed, may reduce any earned patent term adjustment. See 37 CFR 1.704(b). **Status** 1) Responsive to communication(s) filed on Aug 4, 2003 2b) This action is non-final. 2a) This action is FINAL. 3) Since this application is in condition for allowance except for formal matters, prosecution as to the merits is closed in accordance with the practice under Ex parte Quayle, 1935 C.D. 11; 453 O.G. 213. **Disposition of Claims** 4) \bigcirc Claim(s) 56, 60-62, 64, and 65 is/are pending in the application. 4a) Of the above, claim(s) 61 and 62 is/are withdrawn from consideration. 5) Claim(s) is/are allowed. 6) X Claim(s) <u>56, 60, 64, and 65</u> is/are rejected. 7) Claim(s) ______ is/are objected to. 8) Claims are subject to restriction and/or election requirement. **Application Papers** 9) The specification is objected to by the Examiner. 10) ☐ The drawing(s) filed on is/are a) ☐ accepted or b) ☐ objected to by the Examiner. Applicant may not request that any objection to the drawing(s) be held in abeyance. See 37 CFR 1.85(a). 11) ☐ The proposed drawing correction filed on is: a) ☐ approved b) ☐ disapproved by the Examiner. If approved, corrected drawings are required in reply to this Office action. 12) The oath or declaration is objected to by the Examiner. Priority under 35 U.S.C. §§ 119 and 120 13) Acknowledgement is made of a claim for foreign priority under 35 U.S.C. § 119(a)-(d) or (f). a) \square All b) \square Some* c) \square None of: 1. Certified copies of the priority documents have been received. 2. Certified copies of the priority documents have been received in Application No. 3. Copies of the certified copies of the priority documents have been received in this National Stage application from the International Bureau (PCT Rule 17.2(a)). *See the attached detailed Office action for a list of the certified copies not received. 14) Acknowledgement is made of a claim for domestic priority under 35 U.S.C. § 119(e). a) The translation of the foreign language provisional application has been received. 15) ★ Acknowledgement is made of a claim for domestic priority under 35 U.S.C. §§ 120 and/or 121. See Acho. Attachment(s) 1) Notice of References Cited (PTO-892) 4) Interview Summary (PTO-413) Paper No(s). 5) Notice of Informal Patent Application (PTO-152) 2) Notice of Draftsperson's Patent Drawing Review (PTO-948) 6) Other: 3) Information Disclosure Statement(s) (PTO-1449) Paper No(s).

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DETAILED ACTION

- 1. Applicant's Response filed August 4, 2003 (Paper No. 8) is acknowledged. Claims 56, 60 and 64 were amended; claims 1-55, 57-59, 63 and 66-79 were cancelled; and no claims were added. Therefore, claims 56, 60-62, 64 and 65 are currently pending.
- 2. Claims 61 and 62 remain withdrawn from further consideration by the examiner, 37 CFR 1.142(b), as being drawn to non-elected species, there currently being no allowable generic claim.
- 3. Therefore, claims 56, 60, 64 and 65 remain under examination in this action.

Status of Rejections & Objections

4. The previous claim objections are withdrawn in view of applicant's amendments. The rejections under the second paragraph of 35 U.S.C. 112 are also withdrawn in view of applicant's amendments. All other rejections are maintained and applicant's arguments are addressed following each rejection.

Priority

5. Applicant argues in the Response filed August 4, 2003 (page 4) that "there is no requirement that a priority document must disclose or support each and every element of an application or have some exact disclosure relationship as implied by the Examiner".

This is incorrect. As stated previously, applicant has not complied with one or more

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conditions for receiving the benefit of an earlier filing date under 35 U.S.C. 120 and 119(e) specifically because applications 60/030,826, 09/578,282 and 08/969,227 fail to provide adequate support under 35 U.S.C. 112 for the claims of this application since they do not contain a reference to the step of coating a surface with a photobleachable substance as instantly claimed (e.g. step b)). *Elements must have adequate support under 35 U.S.C. 112 in order to enjoy the benefits of priority*. See 35 U.S.C. 120 and 119(e) and also MPEP 201.11. Thus, the denial of the priority claim is maintained and the instant claims are only awarded the date of provisional application 60/233,290, which is August 3, 2000.

Maintained Rejections Claim Rejections - 35 USC § 112

- 6. The following is a quotation of the first paragraph of 35 U.S.C. 112:
 - The specification shall contain a written description of the invention, and of the manner and process of making and using it, in such full, clear, concise, and exact terms as to enable any person skilled in the art to which it pertains, or with which it is most nearly connected, to make and use the same and shall set forth the best mode contemplated by the inventor of carrying out his invention.
- 7. Claims 56, 60, 64 and 65 remain rejected under 35 U.S.C. 112, first paragraph, as containing subject matter which was not described in the specification in such a way as to reasonably convey to one skilled in the relevant art that the inventor(s), at the time the application was filed, had possession of the claimed invention.

To satisfy the written description requirement, an applicant must convey with reasonable clarity to those skilled in the art that, as of the filing date sought, he or she was in possession of the invention. Applicant's claims are directed to a

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"method for synthesizing a polymer array". There are a virtually unlimited number of compounds that would fall within the claimed genus of "polymer array". Moreover, the claims go on to recite that a "photobleachable compound" is used. There are a wide variety of compounds that would fall within this genus as well. The instant specification discloses only very limited examples carrying out the claimed method. This disclosure is neither representative of the claimed genus, nor does it represent a substantial portion of the claimed genus.

Note that the language of the specification should describe the claimed invention so that one skilled in the art can recognize what is claimed. A description of a compound in terms of its function fails to distinguish the compound from others having the same activity or function. A description of what a material does, rather than of what it is, usually does not suffice. The disclosure must allow one skilled in the art to visualize or recognize the identity of the subject matter purportedly described. *University of California v. Eli Lilly and Co.* (U.S. Court of Appeals Federal Circuit (CAFC) 43 USPQ2d 1398 7/22/1997 Decided July 22, 1997; No. 96-1175).

With respect to adequate disclosure of the scope of the presently claimed generic applicant is referred to the discussion in *University of California v. Eli Lilly and Co.* (cited above) regarding disclosure. For adequate disclosure, like enablement, requires representative examples which provide reasonable assurance to one skilled in the art that the compounds falling within the scope both possess the alleged utility and additionally demonstrate that applicant had possession of

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the full scope of the claimed invention. See *In re Riat* (CCPA 1964) 327 F2d 685, 140 USPQ 471; *In re Barr* (CCPA 1971) 444 F 2d 349, 151 USPQ 724 (for enablement) and *University of California v. Eli Lilly and* Co cited above (for disclosure). The more unpredictable the art the greater the showing required (e.g. by "representative examples") for both enablement and adequate disclosure.

Response to Arguments

- 8. Applicant's arguments filed August 4, 2003 have been fully considered but are not found persuasive. The examiner's rationale is set forth below.
- 9. Applicant argues that they are entitled to claims commensurate in scope with the invention and that the claims "should not be limited to mere examples of the invention" (Response, page 5). However, the examiner's position, as stated above, is that the written description requirement is not satisfied since applicant has not conveyed with reasonable clarity to those skilled in the art that, as of the filing date sought, he or she was in possession of the invention.
- 10. As also stated above, applicant's claims are directed to a "method for synthesizing a polymer array". There are a virtually unlimited number of compounds that would fall within the claimed genus of "polymer array". Moreover, the claims go on to recite that a "photobleachable compound" is used. There are a wide variety of compounds that would fall within this genus as well. *The instant specification discloses only very limited*

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examples carrying out the claimed method. This disclosure is neither representative of the claimed genus, nor does it represent a substantial portion of the claimed genus.

- 11. Adequate disclosure requires representative examples. A representative number of examples means that the species that are adequately described are representative of the entire genus. When there is substantial variation within the genus, one must describe a sufficient variety of species to reflect the variation within the genus. Therefore it is deemed that the instant specification lacks adequate support relating to the claimed genus of "polymer array" and "photobleachable compound".
- 12. Note that "the essential goal of the description of the invention requirement is to clearly convey the information that an applicant has invented the subject matter which is claimed." *In re Barker*, 559 F.2d 588, 592 n.4, 194 USPQ 470, 473 n.4 (CCPA 1977), cert. denied, 434 U.S. 1064 (1978). Another objective is to put the public in possession of what the applicant claims as the invention so that the public may ascertain if the patent applicant claims anything that is in common use, or already known. *Evans v. Eaton*, 20 U.S. (7 Wheat.) 356 (1822).
- 13. The examiner deems the art to be unpredictable. The "predictability or lack thereof" in the art refers to the ability of one skilled in the art to extrapolate the disclosed or known results to the claimed invention. If one skilled in the art can readily anticipate the effect of a change within the subject matter to which the claimed invention pertains,

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then there is predictability in the art. On the other hand, if one skilled in the art cannot readily anticipate the effect of a change within the subject matter to which that claimed invention pertains, then there is lack of predictability in the art. Additionally, the Board has held on the issue of unpredictability that "... the unpredictability of an art area alone may be enough to create a reasonable doubt as to the accuracy of statements in the specification." *Ex parte Singh*, 17 U.S.P.Q.2d 1714,1716 (B.P.A.I. 1990).

14. The examiner maintains because of the breadth of the claims, the unpredictability of the art and the lack of representative examples, the written description requirement is not met. Thus the above rejection under 35 U.S.C. 112, first paragraph is maintained.

Maintained Rejections Claim Rejections - 35 USC § 112

15. Claims 56, 60, 64 and 65 remain rejected under 35 U.S.C. 112, first paragraph, because the specification, while being enabling for creation of polynucleotide or polypeptide arrays using pyrylium or diazonium dyes as the photobleachable compound, does not reasonably provide enablement for making **any** "polymer array" using **any** "photobleachable compound or group". The specification does not enable any person skilled in the art to which it pertains, or with which it is most nearly connected, to make and use the invention commensurate in scope with these claims.

It is clear from applicant's specification how one might practice this invention for making polynucleotide or polypeptide arrays using pyrylium or diazonium dyes as the photobleachable compound; however, there is insufficient

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guidance as to how to make/use **any** "polymer array" using **any** "photobleachable compound or group". There are many factors to be considered when determining whether there is sufficient evidence to support a determination that a disclosure does not satisfy the enablement requirement and whether any necessary experimentation is "undue". These factors can include, but are not limited to:

- (1) the breadth of the claims;
- (2) the nature of the invention;
- (3) the state of the prior art;
- (4) the level of one of ordinary skill;
- (5) the level of predictability in the art;
- (6) the amount of direction provided by the inventor;
- (7) the existence of working examples; and
- (8) the quantity of experimentation needed to make or use the invention based on the content of the disclosure.

See In re Wands, 858 F.2d 731, 737, 8 USPQ2d 1400, 1404 (Fed. Cir. 1988).

(1-2) The breadth of the claims and the nature of the invention: The claims are drawn to "method for synthesizing a polymer array" that utilizes a "photobleachable compound or group". Both "polymer array" and "photobleachable compound or group" could read on a wide variety of structures. Such represents broad scope.

(3 and 5) The state of the prior art and the level of predictability in the art:

Various arrays and photobleachable compounds were known in the art at the time of filing; however, only limited numbers of such entities were known and the specification gives no guidance to permit one of skill in the art to devise strategies for the making of *any* "polymer array" and the use of *any* "photobleachable compound or group". The structures of possible variants are sufficiently diverse and one of ordinary skill would not be able to predict their structures. Polymers of various structures would require completely different chemistries. The use of

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different photobleachable compounds or groups would also require differences in preparation and use. One of ordinary skill could not guess, *a priori*, how to make and use **any** such "polymer array[s]" and "photobleachable compound[s] or group[s]" as one could not necessarily predict the required chemistry in the absence of any guidance without undue experimentation. Applicant's claimed scope represents only an invitation to experiment regarding possible "polymer array[s]" and "photobleachable compound[s] or group[s]" of undefined structure.

(4) The level of one of ordinary skill: The level of skill would be high, most likely at the Ph.D. level. However, such persons of ordinary skill in this art, *given its* unpredictability, would have to engage in undue (non-routine) experimentation to carry out the invention as claimed.

- (6-7) The amount of direction provided by the inventor and the existence of working examples: Applicants have provided only very limited examples of the claimed methodology. No generic strategy for determining the required chemistry for making any "polymer array" and using any "photobleachable compound or group" is provided. Specifically, the instant specification fails to identify that structure which is required for the claimed activity. The teachings of the instant specification coupled with the examples only support making polynucleotide or polypeptide arrays using pyrylium or diazonium dyes as the photobleachable compound. See also written description rejection above.
- (8) The quantity of experimentation needed to make or use the invention

 based on the content of the disclosure: The instant specification does not provide

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to one skilled in the art a reasonable amount of guidance with respect to the direction in which the experimentation should proceed in carrying out the full scope of the claimed method. Note that there must be sufficient disclosure, either through illustrative examples or terminology, to teach those of ordinary skill how to make and use the invention as broadly as it is claimed. *In re Vaeck*, 947 F.2d 488, 496 & n.23, 20 USPQ2d 1438, 1445 & n.23 (Fed. Cir. 1991). Therefore, it is deemed that further research of an unpredictable nature would be necessary to make or use the invention as claimed. Thus, due to the inadequacies of the instant disclosure, one of ordinary skill would not have a reasonable expectation of success and the practice of the full scope of the invention would require undue experimentation.

Response to Arguments

- 16. Applicant's arguments filed August 4, 2003 have been fully considered but are not found persuasive. The examiner's rationale is set forth below. Please also see paragraphs 8-14 above.
- 17. Applicant argues (Response, page 6) that the specification "explicitly describes in detail other polymers which may be employed in the context of the instant invention", referring to the application at page 8. While there is a list of different polymers on page 8 of the instant specification, this list in no way enables a method for making arrays of such polymers utilizing the claimed method. There are no examples directed at these other

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types of polymers. As stated in the rejection, the teachings of the instant specification coupled with the examples only support making polynucleotide or polypeptide arrays using pyrylium or diazonium dyes as the photobleachable compound.

- 18. Applicant also argues that "a patent specification needn't set forth every possible embodiment of a generic claim" (Response, pages 6-7). While examples of all embodiments are indeed not required, lack of sufficient working examples, however, is a factor to be considered, especially in a case involving an unpredictable and undeveloped art (see paragraphs 11 & 13 above). One of ordinary skill would not necessarily expect to be able to extrapolate the disclosed examples as far as their applicability to making arrays other than polynucleotide or polypeptide arrays using pyrylium or diazonium dyes as the photobleachable compounds (e.g. making arrays of the different polymers listed on page 8 of the instant specification). In cases involving unpredictable factors, such as most chemical reactions and physiological activity, the scope of enablement obviously varies inversely with the degree of unpredictability of the factors involved. See *In re Fisher*, 57 CCPA 1099,427 F.2d 833,839,166 USPQ 18,24(1970).
- 19. The examiner's position is that the instant specification does not provide to one skilled in the art a reasonable amount of guidance with respect to the direction in which the experimentation should proceed in making and using the claimed invention. In the absence of such guidance, a practitioner of the art would have to resort to a substantial amount of experimental trial and error to predict the required chemistry for making arrays

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other than polynucleotide or polypeptide arrays using pyrylium or diazonium dyes as the photobleachable compounds (e.g. making arrays of the different polymers listed on page 8 of the instant specification). This trial and error would clearly constitute undue experimentation.

20. For these reasons and the reasons of record, the above rejection under 35 U.S.C. 112, first paragraph is maintained.

Maintained Rejections Claim Rejections - 35 USC § 103

- 21. The following is a quotation of 35 U.S.C. 103(a) which forms the basis for all obviousness rejections set forth in this Office action:
 - (a) A patent may not be obtained though the invention is not identically disclosed or described as set forth in section 102 of this title, if the differences between the subject matter sought to be patented and the prior art are such that the subject matter as a whole would have been obvious at the time the invention was made to a person having ordinary skill in the art to which said subject matter pertains. Patentability shall not be negatived by the manner in which the invention was made.
- 22. This application currently names joint inventors. In considering patentability of the claims under 35 U.S.C. 103(a), the examiner presumes that the subject matter of the various claims was commonly owned at the time any inventions covered therein were made absent any evidence to the contrary. Applicant is advised of the obligation under 37 CFR 1.56 to point out the inventor and invention dates of each claim that was not commonly owned at the time a later invention was made in order for the examiner to

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consider the applicability of 35 U.S.C. 103(c) and potential 35 U.S.C. 102(f) or (g) prior art under 35 U.S.C. 103(a).

23. Claims 56, 60, 64 and 65 remain rejected under 35 U.S.C. 103(a) as being unpatentable over Pirrung et al (US 5,143,854), in view of Zebala (US 6,159,681).

Pirrung et al teaches synthesis of polypeptide arrays using photoremovable protecting groups (see Abstract, column 3, Figures 1-7 & 14A). As shown in the reference, polypeptides are built up on a substrate using amino acids with photoremovable protecting groups such as NVOC which reads on the claimed "synthesis intermediates" and "photosensitive protecting groups". The protecting groups are selectively removed by irradiation through a mask; this is performed to create a desired sequence. See, for example, column 28, lines 12-66 of Pirrung et al, which teaches the monomer-by-monomer synthesis of the sequence YGGFL. Pirrung et al teaches that nucleotides can also be used as monomers in their method (see, column 6, lines 9-21, especially line 14), reading on the instant claim 50. The reference also teaches that contrast enhancement materials can be applied between the mask and the substrate to "enhance contrast of light applied to the substrate" (see column 14, lines 5-16). These materials comprise molecules that are "transiently bleached at the wavelength of interest", which allows "greater penetration where light is applied, thereby enhancing contrast" (see column 14, lines 10-16). Molecules such as "quinone diazid" are taught, reading on the instant claims 64 and 65.

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Thus, although Pirrung et al <u>does</u> teach the use of a contrast enhancement layer (as described in the instant specification, page 19, lines 15-23), Pirrung lacks the specific teaching of <u>coating the surface having the synthesis</u> intermediates thereon with a substance comprising a photobleachable compound or group (i.e. instant step b)).

However, using coatings directly on the molecules of interest was well established in the art at the time of filing as taught by Zebala. The reference teachings using a "photoresist layer that is established over a biologic material (which may be immobilized on a substrate). Regions of interest are selected and irradiated to expose specific regions of biologic material" (see Abstract). The biologic materials of Zebala can be peptides or nucleotides (see column 8, lines 46-56). Coating of the photoresist on the biological material is taught, for example, in column 8, line 57 – column 9, line 16 and photoresists are discussed in columns 14-15 of the reference. The reference teaches that using the photoresist directly on the biologic material facilitates analysis of discrete regions (see, e.g. column 6, lines 40-65). Zebala also teaches that contrast enhancement materials can be applied between the mask and the substrate to "enhance contrast of light applied to the photoresist" (see column 31, lines 23-35). These materials comprise molecules that are "transiently bleached by light", which allows "greater penetration where light is applied, thereby enhancing contrast" (see column 31, lines 25-29).

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Therefore, it would have been *prima facie* obvious to one of ordinary skill in the art at the time of filing to carry out Pirrung's synthesis of polypeptide (or polynucleotide) arrays using photoremovable protecting groups further utilizing a photobleachable layer directly coated on the array as taught by Zebala. One would have been motivated to do so due to the advantageous properties of such photobleachable layers (i.e. contrast enhancement materials) as taught by both references, e.g. allowing "greater penetration where light is applied, thereby enhancing contrast". One would have been motivated to place such a layer directly on the materials of interest since such facilitates analysis of discrete regions as taught by Zebala.

Response to Arguments

- 24. Applicant's arguments filed August 4, 2003 have been fully considered but are not found persuasive. The examiner's rationale is set forth below.
- 25. First, applicant argues that Pirrung does not teach steps c), d) or f) of the instant method. The examiner respectfully disagrees. As stated in the rejection, Pirrung clearly contains teachings reading directly on step c). That is, photoremovable protecting groups such as NVOC that are selectively removed by irradiation through a mask are taught and contrast enhancement materials that can be applied between the mask and the substrate to "enhance contrast of light applied to the substrate" (see column 14, lines 5-16) comprising molecules that are "transiently bleached at the wavelength of interest", which

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allows "greater penetration where light is applied, thereby enhancing contrast" (see column 14, lines 10-16) are also taught. Step d) of removing the substance comprising a photobleachable compound would be inherently present in the method of Pirrung during the creation of the array.

- 26. The examiner noted in the rejection that Pirrung lacks the specific teaching of coating the surface having the synthesis intermediates thereon with a substance comprising a photobleachable compound or group (i.e. instant step b)). However, the examiner's position is that this feature is taught by Zebala, as further discussed below. The "recoating" step f) is an optional step ("if another synthesis intermediate is to be added") and thus is not afforded any patentable weight. Applicant's arguments directed to the Zebala reference not teaching step f) are not persuasive for the same reasons.
- 27. Next, applicant also refers on pages 8 and 9 of the Response to step c) of their method being a step of "irradiating the surface and simultaneously removing both the photobleachable compound and the photoprotective group" (e.g. see page 9, 1st full paragraph). Step c) does *not* contain such a recitation. The step recites that at least a part of the surface is irradiated to "bleach said photobleachable compound" and "remove said protective group". There is nothing in step c) regarding *removing* the photobleachable compound.

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- 28. Also, with respect to applicants further arguments concerning the Zebala reference, applicants argue that "there is no teaching or suggestion that the photoresist layer may be used in the construction of an array of polymers" (Response, page 9). However, Pirrung clearly teaches synthesis of polypeptide arrays using photoremovable protecting groups and contrast enhancement materials and it is the *combination* of references that is being relied upon. Note that test for obviousness is not whether the features of a secondary reference may be bodily incorporated into the structure of the primary reference; nor is it that the claimed invention must be expressly suggested in any one or all of the references. Rather, the test is what the combined teachings of the references would have suggested to those of ordinary skill in the art. See *In re Keller*, 642 F.2d 413, 208 USPQ 871 (CCPA 1981).
- 29. The examiner recognizes that obviousness can only be established by combining or modifying the teachings of the prior art to produce the claimed invention where there is some teaching, suggestion, or motivation to do so found either in the references themselves or in the knowledge generally available to one of ordinary skill in the art. See *In re Fine*, 837 F.2d 1071, 5 USPQ2d 1596 (Fed. Cir. 1988) and *In re Jones*, 958 F.2d 347, 21 USPQ2d 1941 (Fed. Cir. 1992).
- 30. In this case, the examiner maintains that the *combined* teachings of the cited references render the claimed invention obvious. The strongest rationale for combining references is a recognition, expressly or impliedly in the prior art or drawn from a

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convincing line of reasoning based on established scientific principles or legal precedent, that some advantage or expected beneficial result would have been produced by their combination. *In re Sernaker*, 702 F.2d 989, 994-95, 217 USPQ 1, 5-6 (Fed. Cir. 1983). In the instant case, the beneficial result of placing a photobleachable layer directly on the array is to facilitate analysis of discrete regions.

31. For these reasons and the reasons of record, the above rejection under 35 U.S.C. 103 is maintained.

Status of Claims/Conclusion

- 32. No claims are allowed.
- 33. THIS ACTION IS MADE FINAL. Applicant is reminded of the extension of time policy as set forth in 37 CFR 1.136(a).

A shortened statutory period for reply to this final action is set to expire THREE MONTHS from the mailing date of this action. In the event a first reply is filed within TWO MONTHS of the mailing date of this final action and the advisory action is not mailed until after the end of the THREE-MONTH shortened statutory period, then the shortened statutory period will expire on the date the advisory action is mailed, and any extension fee pursuant to 37 CFR 1.136(a) will be calculated from the mailing date of the advisory action. In no

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event, however, will the statutory period for reply expire later than SIX MONTHS from the mailing date of this final action.

34. Any inquiry concerning this communication or earlier communications from the

examiner should be directed to Maurie Garcia Baker, Ph.D. whose telephone number is

(703) 308-0065. The examiner is on an increased flextime schedule but can normally be

reached on Monday-Thursday and alternate Fridays from 9:30 to 7:00.

35. If attempts to reach the examiner by telephone are unsuccessful, the examiner's

supervisor, Andrew J. Wang, can be reached at (703) 306-3217. The fax phone number

for the organization where this application or proceeding is assigned is (703) 872-9306.

Any inquiry of a general nature or relating to the status of this application or proceeding

should be directed to the receptionist whose telephone number is (703) 308-0196.

Maurie Garcia Baker, Ph.D. October 17, 2003

MAURIE GARCIA BAIGER PH.D